



SEQUENCE LISTING

b1  
<110> Imamura, Toru

Asada, Masahiro

Oka, Syuichi

Suzuki, Masashi

Yoneda, Atsuko

Ota, Keiko

Oda, Yuko

Miyakawa, Kazuko

Orikasa, Noriko

Asada, Chie

Kojima, Tetsuhito

<120> HEPARIN-BINDING PROTEINS MODIFIED WITH SUGAR CHAINS,  
METHOD OF PRODUCING THE SAME AND PHARMACEUTICAL  
COMPOSITIONS CONTAINING THE SAME

<130> 382.1019

<140> 09/121,017

<141> 1998-07-22

<150> 307721/1997

<151> 1997-11-10

<160> 31

<170> PatentIn Ver. 2.0

<210> 1

<211> 221

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human rydolan and a part of human fibroblast

growth factor 1

<400> 1

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35 40 45

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50 55 60

Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His

65 70 75 80

Pro Leu Val Pro Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr

85

*Start of human fibroblast growth factor*

90

95

Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val

100 105 110

Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser  
115 120 125

Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln  
130 135 140

Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro  
145 150 155 160

Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn  
165 170 175

Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu  
180 185 190

Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln  
195 200 205

Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp  
210 215 220

<210> 2

<211> 663

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human rydocan and a part of human fibroblast

growth factor 1

<220>

<221> CDS

<222> (1)..(663)

<400> 2

atg gcc ccc gcc cgt ctg ttc gcg ctg ctg ctg ttc ttc gta ggc gga 48

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

gtc gcc gag tcg atc cga gag act gag gtc atc gac ccc cag gac ctc 96

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

cta gaa ggc cga tac ttc tcc gga gcc cta cca gac gat gag gat gta 144

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35 40 45

gtg ggg ccc ggg cag gaa tct gat gac ttt gag ctg tct ggc tct gga 192

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50 55 60

gat ctg gat gac ttg gaa gac tcc atg atc ggc cct gaa gtt gtc cat 240

Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His

65 70 75 80

ccc ttg gtg cct cta gat gct aat tac aag aag ccc aaa ctc ctc tac 288

Pro Leu Val Pro Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr

| 85  | 90  | 95  |     |
|---|-----|-----|-----|
| tgt agc aac ggg ggc cac ttc ctg agg atc ctt ccg gat ggc aca gtg |     |     | 336 |
| Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val |     |     |     |
| 100   | 105 | 110 |     |
|   |     |     |     |
| gat ggg aca agg gac agg agc gac cag cac att cag ctg cag ctc agt |     |     | 384 |
| Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser |     |     |     |
| 115   | 120 | 125 |     |
|   |     |     |     |
| gcg gaa agc gtg ggg gag gtg tat ata aag agt acc gag act ggc cag |     |     | 432 |
| Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln |     |     |     |
| 130   | 135 | 140 |     |
|   |     |     |     |
| tac ttg gcc atg gac acc gac ggg ctt tta tac ggc tca cag aca cca |     |     | 480 |
| Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro |     |     |     |
| 145   | 150 | 155 | 160 |
|   |     |     |     |
| aat gag gaa tgt ttg ttc ctg gaa agg ctg gag gag aac cat tac aac |     |     | 528 |
| Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn |     |     |     |
| 165   | 170 | 175 |     |
|   |     |     |     |
| acc tat ata tcc aag aag cat gca gag aag aat tgg ttt gtt ggc ctc |     |     | 576 |
| Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu |     |     |     |
| 180   | 185 | 190 |     |
|   |     |     |     |
| aag aag aat ggg agc tgc aaa cgc ggt cct cgg act cac tat ggc cag |     |     | 624 |
| Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln |     |     |     |
| 195   | 200 | 205 |     |
|   |     |     |     |
| aaa gca atc ttg ttt ctc ccc ctg cca gtc tct tct gat             |     |     | 663 |

Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp

210

215

220

<210> 3

<211> 175

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of mouse fibroblast growth factor 6 and

a part of human fibroblast growth factor 1

<400> 3

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1

5

10

15

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20

25

30

Arg Ala Asn Gly Thr Leu Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu.

35

*Start<sup>40</sup> of human fibroblast growth factor<sup>45</sup>*

Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly

50

55

60

Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln

65

70

75

80

Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr

85

90

95

Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln

100

105

110

Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His

115

120

125

Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val

130

135

140

Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr

145

150

155

160

Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp

165

170

175

<210> 4

<211> 525

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
sequence for a part of mouse fibroblast growth factor 6 and  
a part of human fibroblast growth factor 1

<220>

<221> CDS

<222> (1)..(525)

*end of human  
fibroblast  
growth  
factor 1*

<400> 4

atg tcc cgg gga gca gga cgt gtt cag ggc acg ctg cag gct ctc gtc 48

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1 5 10 15

ttc tta ggc gtc cta gtg ggc atg gtg gtg ccc tca ~~cct~~ gcc ggc gcc 96

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20 25 30

cgc gcc aac ggc acg cta ctg gac gct aat tac aag aag ccc aaa ctc 144

Arg Ala Asn Gly Thr Leu Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu

35 40 45

ctc tac tgt agc aac ggg ggc cac ttc ctg agg atc ctt ccg gat ggc 192

Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly

50 55 60

aca gtg gat ggg aca agg gac agg agc gac cag cac att cag ctg cag 240

Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln

65 70 75 80

ctc agt gcg gaa agc gtg ggg gag gtg tat ata aag agt acc gag act 288

Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr

85 90 95

ggc cag tac ttg gcc atg gac acc gac ggg ctt tta tac ggc tca cag 336

Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln

100 105 110



aca cca aat gag gaa tgt ttg ttc ctg gaa agg ctg gag gag aac cat 384  
 Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His  
 115 120 125

tac aac acc tat ata tcc aag aag cat gca gag aag aat tgg ttt gtt 432  
 Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val  
 130 135 140

ggc ctc aag aag aat ggg agc tgc aaa cgc ggt cct cgg act cac tat 480  
 Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr  
 145 150 155 160

ggc cag aaa gca atc ttg ttt ctc ccc ctg cca gtc tct tct gat 525  
 Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp  
 165 170 175

<210> 5

<211> 181

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
 sequence for a part of mouse fibroblast growth factor 6,  
 a part of human fibroblast growth factor 1 and an artificial  
 sequence

<400> 5

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

|     |     |     |     |
|-----|-----|-----|-----|
| 1   | 5   | 10  | 15  |
| Phe | Leu | Gly | Val |
| Leu | Val | Gly | Met |
| Val | Val | Pro | Ser |
| Pro | Ala | Gly | Ala |
| 20  | 25  | 30  |     |
| Arg | Ala | Gln | Gly |
| Thr | Leu | Leu | Asp |
| Ala | Asn | Tyr | Lys |
| Lys | Lys | Pro | Lys |
| Leu |     |     |     |
| 35  | 40  | 45  |     |
| Leu | Tyr | Cys | Ser |
| Asn | Gly | Gly | His |
| Phe | Leu | Arg | Ile |
| Leu | Pro | Asp | Gly |
| 50  | 55  | 60  |     |
| Thr | Val | Asp | Gly |
| Thr | Arg | Asp | Arg |
| Ser | Asp | Gln | His |
| Ile | Gln | Leu | Gln |
| 65  | 70  | 75  | 80  |
| Leu | Ser | Ala | Glu |
| Ser | Val | Gly | Glu |
| Val | Tyr | Ile | Lys |
| Ser | Thr | Glu | Thr |
| 85  | 90  | 95  |     |
| Gly | Gln | Tyr | Leu |
| Ala | Met | Asp | Thr |
| Asp | Gly | Leu | Leu |
| Tyr | Gly | Ser | Gln |
| 100 | 105 | 110 |     |
| Thr | Pro | Asn | Glu |
| Glu | Cys | Leu | Phe |
| Leu | Glu | Arg | Leu |
| Glu | Glu | Ala | Ala |
| 115 | 120 | 125 |     |
| Thr | Pro | Ala | Pro |
| Asn | His | Tyr | Asn |
| Thr | Tyr | Ile | Ser |
| Lys | Lys | His | Ala |
| 130 | 135 | 140 |     |
| Glu | Lys | Asn | Trp |
| Phe | Val | Gly | Leu |
| Lys | Lys | Asn | Gly |
| Ser | Cys | Lys | Arg |
| 145 | 150 | 155 | 160 |
| Gly | Pro | Arg | Thr |
| His | Tyr | Gly | Gln |
| Lys | Ala | Ile | Leu |
| Phe | Leu | Pro | Leu |

*start of human fibroblast growth factor 1*

*inserted  
artificial  
sequence*

165

170

175

Pro Val Ser Ser Asp

180

&lt;210&gt; 6

&lt;211&gt; 543

&lt;212&gt; DNA

&lt;213&gt; Artificial Sequence

&lt;220&gt;

&lt;223&gt; Description of Artificial Sequence: fusion of

sequence for a part of mouse fibroblast growth factor 6,

a part of human fibroblast growth factor 1 and an artificial

sequence

&lt;220&gt;

&lt;221&gt; CDS

&lt;222&gt; (1)..(543)

&lt;400&gt; 6

atg tcc cgg gga gca gga cgt gtt cag ggc acg ctg cag gct ctc gtc 48

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1

5

10

15

ttc tta ggc gtc cta gtg ggc atg gtg gtg ccc tca cct gcc ggc gcc 96

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20

25

30

cgc gcc caa ggc acg cta ctg gac gct aat tac aag aag ccc aaa ctc 144

|     |     |     |     |     |     |     |     |     |     |     |     |     |     |     |     |     |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Arg | Ala | Gln | Gly | Thr | Leu | Leu | Asp | Ala | Asn | Tyr | Lys | Lys | Pro | Lys | Leu |     |
|     |     | 35  |     |     |     |     | 40  |     |     |     |     | 45  |     |     |     |     |
| ctc | tac | tgt | agc | aac | ggg | ggc | cac | ttc | ctg | agg | atc | ctt | ccg | gat | ggc | 192 |
| Leu | Tyr | Cys | Ser | Asn | Gly | Gly | His | Phe | Leu | Arg | Ile | Leu | Pro | Asp | Gly |     |
|     |     | 50  |     |     |     |     | 55  |     |     |     |     | 60  |     |     |     |     |
| aca | gtg | gat | ggg | aca | agg | gac | agg | agc | gac | cag | cac | att | cag | ctg | cag | 240 |
| Thr | Val | Asp | Gly | Thr | Arg | Asp | Arg | Ser | Asp | Gln | His | Ile | Gln | Leu | Gln |     |
|     |     | 65  |     |     |     | 70  |     |     |     |     | 75  |     |     |     | 80  |     |
| ctc | agt | gcg | gaa | agc | gtg | ggg | gag | gtg | tat | ata | aag | agt | acc | gag | act | 288 |
| Leu | Ser | Ala | Glu | Ser | Val | Gly | Glu | Val | Tyr | Ile | Lys | Ser | Thr | Glu | Thr |     |
|     |     |     |     |     | 85  |     |     |     |     | 90  |     |     |     | 95  |     |     |
| ggc | cag | tac | ttg | gcc | atg | gac | acc | gac | ggg | ctt | tta | tac | ggc | tca | cag | 336 |
| Gly | Gln | Tyr | Leu | Ala | Met | Asp | Thr | Asp | Gly | Leu | Leu | Tyr | Gly | Ser | Gln |     |
|     |     |     | 100 |     |     |     |     | 105 |     |     |     |     | 110 |     |     |     |
| aca | cca | aat | gag | gaa | tgt | ttg | ttc | ctg | gaa | agg | ctg | gag | gag | gct | gct | 384 |
| Thr | Pro | Asn | Glu | Glu | Cys | Leu | Phe | Leu | Glu | Arg | Leu | Glu | Glu | Ala | Ala |     |
|     |     | 115 |     |     |     |     |     | 120 |     |     |     |     | 125 |     |     |     |
| act | cca | gct | cca | aac | cat | tac | aac | acc | tat | ata | tcc | aag | aag | cat | gca | 432 |
| Thr | Pro | Ala | Pro | Asn | His | Tyr | Asn | Thr | Tyr | Ile | Ser | Lys | Lys | His | Ala |     |
|     |     | 130 |     |     |     |     | 135 |     |     |     |     | 140 |     |     |     |     |
| gag | aag | aat | tgg | ttt | gtt | ggc | ctc | aag | aag | aat | ggg | agc | tgc | aaa | cgc | 480 |
| Glu | Lys | Asn | Trp | Phe | Val | Gly | Leu | Lys | Lys | Asn | Gly | Ser | Cys | Lys | Arg |     |
| 145 |     |     |     |     |     | 150 |     |     |     | 155 |     |     |     |     | 160 |     |

ggt cct cgg act cac tat ggc cag aaa gca atc ttg ttt ctc ccc ctg 528

Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu

165

170

175

cca gtc tct tct gat

543

Pro Val Ser Ser Asp

180

<210> 7

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 7

ttgtcgaccc accatggccc ccgcccgtct

30

<210> 8

<211> 26

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 8

ttgatatcta gaggcaccaa gggatg

26

<210> 9

<211> 35

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 9

gcgtcgacag cgctaattac aagaagccca aactc

35

<210> 10

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 10

ccgaattcga attctttaat cagaagagac tgg

33

<210> 11

<211> 64

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 11

gcgtcgaccc accatgtccc ggggagcagg acgtgttcag ggcacgctgc aggctctcgt 60

cttc

64

<210> 12

<211> 29

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 12

gcgatatcca gtagcgtgcc gttggcgcg

29

<210> 13

<211> 18

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 13

gcgtcgaccc accatgtc

18

<210> 14

<211> 29

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 14

gcgatatcca gtagcgtgcc ttgggcgcg

29

<210> 15

<211> 38

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 15

gctggaggag gctgctactc cagctccaaa ccattaca

38

<210> 16

<211> 21



<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 16

gccgctctag aactagtgga t

21

<210> 17

<211> 200

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human  
fibroblast

growth factor 1

<400> 17

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1

5

10

15

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20

25

30

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35

40

45

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50

55

60

Asp Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys Ser Asn Gly Gly

65

70

75

80

*start of FGF1*

His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp Gly Thr Arg Asp

85

90

95

Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala Glu Ser Val Gly

100

105

110

Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr Leu Ala Met Asp

115

120

125

Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn Glu Glu Cys Leu

130

135

140

Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr Tyr Ile Ser Lys

145

150

155

160

Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly Ser

165

170

175

Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe

180

185

190

Leu Pro Leu Pro Val Ser Ser Asp

195

200

<210> 18

<211> 600

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human fibroblast

growth factor 1

<220>

<221> CDS

<222> (1)..(600)

<400> 18

atg gcc ccc gcc cgt ctg ttc gcg ctg ctg ctg ttc ttc gta ggc gga 48

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

gtc gcc gag tcg atc cga gag act gag gtc atc gac ccc cag gac ctc 96

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

cta gaa ggc cga tac ttc tcc gga gcc cta cca gac gat gag gat gta 144

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35 40 45

gtg ggg ccc ggg cag gaa tct gat gac ttt gag ctg tct ggc tct gga 192

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50 55 60



Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe

180

185

190

ctc ccc ctg cca gtc tct tct gat

600

Leu Pro Leu Pro Val Ser Ser Asp

195

200

<210> 19

<211> 200

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
sequence for a part of human ryudocan mutant and a part of human  
fibroblast growth factor 1

<400> 19

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1

5

10

15

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20

25

30

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Ser Asp Asp Glu Asp Val

35

40

45

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50

55

60

Asp Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys Ser Asn Gly Gly  
65 70 75 80

His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp Gly Thr Arg Asp  
85 90 95

Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala Glu Ser Val Gly  
100 105 110

Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr Leu Ala Met Asp  
115 120 125

Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn Glu Glu Cys Leu  
130 135 140

Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr Tyr Ile Ser Lys  
145 150 155 160

Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly Ser  
165 170 175

Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe  
180 185 190

Leu Pro Leu Pro Val Ser Ser Asp  
195 200

<210> 20

<211> 600

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
sequence for a part of human ryudocan mutant and a part of human  
fibroblast growth factor 1

<220>

<221> CDS

<222> (1)..(600)

<400> 20

atg gcc ccc gcc cgt ctg ttc gcg ctg ctg ctg ttc ttc gta ggc gga 48

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

gtc gcc gag tcg atc cga gag act gag gtc atc gac ccc cag gac ctc 96

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

cta gaa ggc cga tac ttc tcc gga gcc cta tca gac gat gag gat gta 144

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Ser Asp Asp Glu Asp Val

35 40 45

gtg ggg ccc ggg cag gaa tct gat gac ttt gag ctg tct ggc tct gga 192

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50 55 60

gat gct aat tac aag aag ccc aaa ctc ctc tac tgt agc aac ggg ggc 240

Asp Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys Ser Asn Gly Gly

| 65  | 70  | 75  | 80  |     |
|---|-----|-----|-----|-----|
| cac ttc ctg agg atc ctt ccg gat ggc aca gtg gat ggg aca agg gac |     |     |     | 288 |
| His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp Gly Thr Arg Asp |     |     |     |     |
|   | 85  | 90  | 95  |     |
| agg agc gac cag cac att cag ctg cag ctc agt gcg gaa agc gtg ggg |     |     |     | 336 |
| Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala Glu Ser Val Gly |     |     |     |     |
|   | 100 | 105 | 110 |     |
| gag gtg tat ata aag agt acc gag act ggc cag tac ttg gcc atg gac |     |     |     | 384 |
| Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr Leu Ala Met Asp |     |     |     |     |
|   | 115 | 120 | 125 |     |
| acc gac ggg ctt tta tac ggc tca cag aca cca aat gag gaa tgt ttg |     |     |     | 432 |
| Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn Glu Glu Cys Leu |     |     |     |     |
|   | 130 | 135 | 140 |     |
| ttc ctg gaa agg ctg gag gag aac cat tac aac acc tat ata tcc aag |     |     |     | 480 |
| Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr Tyr Ile Ser Lys |     |     |     |     |
|   | 145 | 150 | 155 | 160 |
| aag cat gca gag aag aat tgg ttt gtt ggc ctc aag aag aat ggg agc |     |     |     | 528 |
| Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly Ser |     |     |     |     |
|   | 165 | 170 | 175 |     |
| tgc aaa cgc ggt cct cgg act cac tat ggc cag aaa gca atc ttg ttt |     |     |     | 576 |
| Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe |     |     |     |     |
|   | 180 | 185 | 190 |     |



ctc ccc ctg cca gtc tct tct gat

600

Leu Pro Leu Pro Val Ser Ser Asp

195

200

<210> 21

<211> 254

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human  
fibroblast

growth factor 1

<400> 21

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1

5

10

15

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20

25

30

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35

40

45

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50

55

60

Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His

52

|     |     |     |     |     |                      |     |     |     |     |     |     |     |     |     |     |
|-----|-----|-----|-----|-----|----------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 65  |     | 70  |     | 75  |                      | 80  |     |     |     |     |     |     |     |     |     |
| Pro | Leu | Val | Pro | Leu | Asp                  | Asn | His | Ile | Pro | Glu | Arg | Ala | Gly | Ser | Gly |
|     |     |     | 85  |     |                      |     |     |     | 90  |     |     |     |     | 95  |     |
| Ser | Gln | Val | Pro | Thr | Glu                  | Pro | Lys | Lys | Leu | Glu | Glu | Asn | Glu | Val | Ile |
|     |     | 100 |     |     |                      |     |     | 105 |     |     |     |     | 110 |     |     |
| Pro | Lys | Arg | Ile | Ser | Pro                  | Val | Ala | Asn | Tyr | Lys | Lys | Pro | Lys | Leu | Leu |
|     | 115 |     |     |     |                      | 120 |     |     |     |     |     |     |     | 125 |     |
|     |     |     |     |     | <i>start of FGF1</i> |     |     |     |     |     |     |     |     |     |     |
| Tyr | Cys | Ser | Asn | Gly |                      |     |     |     |     | Gly | His | Phe | Leu | Arg | Ile |
|     | 130 |     |     |     |                      | 135 |     |     |     |     | 140 |     |     |     |     |
| Val | Asp | Gly | Thr | Arg | Asp                  | Arg | Ser | Asp | Gln | His | Ile | Gln | Leu | Gln | Leu |
| 145 |     |     |     |     | 150                  |     |     |     |     | 155 |     |     | 160 |     |     |
| Ser | Ala | Glu | Ser | Val | Gly                  | Glu | Val | Tyr | Ile | Lys | Ser | Thr | Glu | Thr | Gly |
|     |     |     | 165 |     |                      |     |     |     | 170 |     |     |     | 175 |     |     |
| Gln | Tyr | Leu | Ala | Met | Asp                  | Thr | Asp | Gly | Leu | Leu | Tyr | Gly | Ser | Gln | Thr |
|     |     | 180 |     |     |                      |     |     | 185 |     |     |     |     | 190 |     |     |
| Pro | Asn | Glu | Glu | Cys | Leu                  | Phe | Leu | Glu | Arg | Leu | Glu | Glu | Asn | His | Tyr |
|     | 195 |     |     |     |                      | 200 |     |     |     |     |     |     | 205 |     |     |
| Asn | Thr | Tyr | Ile | Ser | Lys                  | Lys | His | Ala | Glu | Lys | Asn | Trp | Phe | Val | Gly |
|     | 210 |     |     |     |                      | 215 |     |     |     |     | 220 |     |     |     |     |
| Leu | Lys | Lys | Asn | Gly | Ser                  | Cys | Lys | Arg | Gly | Pro | Arg | Thr | His | Tyr | Gly |
| 225 |     |     |     | 230 |                      |     |     |     | 235 |     |     |     | 240 |     |     |

Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp

245

250

<210> 22

<211> 762

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human  
fibroblast

growth factor 1

<220>

<221> CDS

<222> (1)..(762)

<400> 22

atg gcc ccc gcc cgt ctg ttc gcg ctg ctg ctg ttc ttc gta ggc gga 48

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

gtc gcc gag tcg atc cga gag act gag gtc atc gac ccc cag gac ctc 96

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

cta gaa ggc cga tac ttc tcc gga gcc cta cca gac gat gag gat gta 144

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

|  |     |     |     |
|--|-----|-----|-----|
| 35   | 40  | 45  |     |
| gtg ggg ccc ggg cag gaa tct gat gac ttt gag ctg tct ggc tct gga<br>Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly |     |     | 192 |
| 50   | 55  | 60  |     |
| gat ctg gat gac ttg gaa gac tcc atg atc ggc cct gaa gtt gtc cat<br>Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His |     |     | 240 |
| 65   | 70  | 75  | 80  |
| ccc ttg gtg cct cta gat aac cat atc cct gag agg gca ggg tct ggg<br>Pro Leu Val Pro Leu Asp Asn His Ile Pro Glu Arg Ala Gly Ser Gly |     |     | 288 |
|  | 85  | 90  | 95  |
| agc caa gtc ccc acc gaa ccc aag aaa cta gag gag aat gag gtt atc<br>Ser Gln Val Pro Thr Glu Pro Lys Lys Leu Glu Glu Asn Glu Val Ile |     |     | 336 |
|  | 100 | 105 | 110 |
| ccc aag aga atc tca ccc gtt gct aat tac aag aag ccc aaa ctc ctc<br>Pro Lys Arg Ile Ser Pro Val Ala Asn Tyr Lys Lys Pro Lys Leu Leu |     |     | 384 |
|  | 115 | 120 | 125 |
| tac tgt agc aac ggg ggc cac ttc ctg agg atc ctt ccg gat ggc aca<br>Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr |     |     | 432 |
|  | 130 | 135 | 140 |
| gtg gat ggg aca agg gac agg agc gac cag cac att cag ctg cag ctc<br>Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu |     |     | 480 |
| 145  | 150 | 155 | 160 |

agc ggc gaa agc gtc ggg gag gtc tat ata aag agc acc gag act ggc 528  
 Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly  
 165 170 175

cag tac ttg gcc atg gac acc gac ggg ctt tta tac ggc tca cag aca 576  
 Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr  
 180 185 190

cca aat gag gaa tgt ttg ttc ctg gaa agg ctg gag gag aac cat tac 624  
 Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr  
 195 200 205

aac acc tat ata tcc aag aag cat gca gag aag aat tgg ttt gtt ggc 672  
 Asn Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly  
 210 215 220

ctc aag aag aat ggg agc tgc aaa cgc ggt cct cgg act cac tat ggc 720  
 Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly  
 225 230 235 240

cag aaa gca atc ttg ttt ctc ccc ctg cca gtc tct tct gat 762  
 Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp  
 245 250

<210> 23

<211> 281

<212> PRT

<213> Artificial Sequence

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human fibroblast

growth factor 1

<400> 23

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly  
1 5 10 15

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu  
20 25 30

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val  
35 40 45

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly  
50 55 60

Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His  
65 70 75 80

Pro Leu Val Pro Leu Asp Asn His Ile Pro Glu Arg Ala Gly Ser Gly  
85 90 95

Ser Gln Val Pro Thr Glu Pro Lys Lys Leu Glu Glu Asn Glu Val Ile  
100 105 110

Pro Lys Arg Ile Ser Pro Val Glu Glu Ser Glu Asp Val Ser Asn Lys  
115 120 125

Val Ser Met Ser Ser Thr Val Gln Gly Ser Asn Ile Phe Glu Arg Thr  
130 135 140

human  
 human  
 HGF-1

Glu Val Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys Ser Asn Gly  
 145 150 155 160

Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp Gly Thr Arg  
 165 170 175

Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala Glu Ser Val  
 180 185 190

Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr Leu Ala Met  
 195 200 205

Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn Glu Glu Cys  
 210 215 220

Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr Tyr Ile Ser  
 225 230 235 240

Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly  
 245 250 255

Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu  
 260 265 270

Phe Leu Pro Leu Pro Val Ser Ser Asp  
 275 280

<210> 24

<211> 843

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of human ryudocan and a part of human  
fibroblast

growth factor 1

<220>

<221> CDS

<222> (1)..(843)

<400> 24

atg gcc ccc gcc cgt ctg ttc gcg ctg ctg ctg ttc ttc gta ggc gga 48

Met Ala Pro Ala Arg Leu Phe Ala Leu Leu Leu Phe Phe Val Gly Gly

1 5 10 15

gtc gcc gag tcg atc cga gag act gag gtc atc gac ccc cag gac ctc 96

Val Ala Glu Ser Ile Arg Glu Thr Glu Val Ile Asp Pro Gln Asp Leu

20 25 30

cta gaa ggc cga tac ttc tcc gga gcc cta cca gac gat gag gat gta 144

Leu Glu Gly Arg Tyr Phe Ser Gly Ala Leu Pro Asp Asp Glu Asp Val

35 40 45

gtg ggg ccc ggg cag gaa tct gat gac ttt gag ctg tct ggc tct gga 192

Val Gly Pro Gly Gln Glu Ser Asp Asp Phe Glu Leu Ser Gly Ser Gly

50 55 60



|  |     |
|--|-----|
| gat ctg gat gac ttg gaa gac tcc atg atc ggc cct gaa gtt gtc cat  | 240 |
| Asp Leu Asp Asp Leu Glu Asp Ser Met Ile Gly Pro Glu Val Val His  |     |
| 65                                      70                                      75                                      80     |     |
|  |     |
| ccc ttg gtg cct cta gat aac cat atc cct gag agg gca ggg tct ggg  | 288 |
| Pro Leu Val Pro Leu Asp Asn His Ile Pro Glu Arg Ala Gly Ser Gly  |     |
| 85                                      90                                      95   |     |
|  |     |
| agc caa gtc ccc acc gaa ccc aag aaa cta gag gag aat gag gtt atc  | 336 |
| Ser Gln Val Pro Thr Glu Pro Lys Lys Leu Glu Glu Asn Glu Val Ile  |     |
| 100                                      105                                      110  |     |
|  |     |
| ccc aag aga atc tca ccc gtt gaa gag agt gag gat gtg tcc aac aag  | 384 |
| Pro Lys Arg Ile Ser Pro Val Glu Glu Ser Glu Asp Val Ser Asn Lys  |     |
| 115                                      120                                      125  |     |
|  |     |
| gtg tca atg tcc agc act gtg cag ggc agc aac atc ttt gag aga acg  | 432 |
| Val Ser Met Ser Ser Thr Val Gln Gly Ser Asn Ile Phe Glu Arg Thr  |     |
| 130                                      135                                      140  |     |
|  |     |
| gag gtc gct aat tac aag aag ccc aaa ctc ctc tac tgt agc aac ggg  | 480 |
| Glu Val Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys Ser Asn Gly  |     |
| 145                                      150                                      155                                      160 |     |
|  |     |
| ggc cac ttc ctg agg atc ctt ccg gat ggc aca gtg gat ggg aca agg  | 528 |
| Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp Gly Thr Arg  |     |
| 165                                      170                                      175  |     |
|  |     |
| gac agg agc gac cag cac att cag ctg cag ctc agt gcg gaa agc gtg  | 576 |
| Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala Glu Ser Val  |     |
| 180                                      185                                      190  |     |

ggg gag gtg tat ata aag agt acc gag act ggc cag tac ttg gcc atg 624  
 Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr Leu Ala Met

195 200 205

gac acc gac ggg ctt tta tac ggc tca cag aca cca aat gag gaa tgt 672  
 Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn Glu Glu Cys

210 215 220

ttg ttc ctg gaa agg ctg gag gag aac cat tac aac acc tat ata tcc 720  
 Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr Tyr Ile Ser

225 230 235 240

aag aag cat gca gag aag aat tgg ttt gtt ggc ctc aag aag aat ggg 768  
 Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly

245 250 255

agc tgc aaa cgc ggt cct cgg act cac tat ggc cag aaa gca atc ttg 816  
 Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu

260 265 270

ttt ctc ccc ctg cca gtc tct tct gat 843  
 Phe Leu Pro Leu Pro Val Ser Ser Asp

275 280

<210> 25

<211> 172

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of mouse fibroblast growth factor 6 and  
a part of human fibroblast growth factor 1

<400> 25

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val  
1 5 10 15

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala  
20 25 30

Arg Ala Asn Gly Ser Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys  
35 40 45

Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp  
50 55 60

Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala  
65 70 75 80

Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr  
85 90 95

Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn  
100 105 110

Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr  
115 120 125

Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys  
 130 135 140

Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys  
 145 150 155 160

Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp  
 165 170

<210> 26

<211> 516

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
 sequence for a part of mouse fibroblast growth factor 6 and  
 a part of human fibroblast growth factor 1

<220>

<221> CDS

<222> (1)..(516)

<400> 26

atg tcc cgg gga gca gga cgt gtt cag ggc acg ctg cag gct ctc gtc 48  
 Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val  
 1 5 10 15

ttc tta ggc gtc cta gtg ggc atg gtg gtg ccc tca cct gcc ggc gcc 96

|   |     |
|---|-----|
| Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala |     |
| 20 25 30  |     |
| cgc gcc aac ggc tcg gct aat tac aag aag ccc aaa ctc ctc tac tgt | 144 |
| Arg Ala Asn Gly Ser Ala Asn Tyr Lys Lys Pro Lys Leu Leu Tyr Cys |     |
| 35 40 45  |     |
| agc aac ggg ggc cac ttc ctg agg atc ctt ccg gat ggc aca gtg gat | 192 |
| Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly Thr Val Asp |     |
| 50 55 60  |     |
| ggg aca agg gac agg agc gac cag cac att cag ctg cag ctc agt gcg | 240 |
| Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln Leu Ser Ala |     |
| 65 70 75 80   |     |
| gaa agc gtg ggg gag gtg tat ata aag agt acc gag act ggc cag tac | 288 |
| Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr Gly Gln Tyr |     |
| 85 90 95  |     |
| ttg gcc atg gac acc gac ggg ctt tta tac ggc tca cag aca cca aat | 336 |
| Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln Thr Pro Asn |     |
| 100 105 110   |     |
| gag gaa tgt ttg ttc ctg gaa agg ctg gag gag aac cat tac aac acc | 384 |
| Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn His Tyr Asn Thr |     |
| 115 120 125   |     |
| tat ata tcc aag aag cat gca gag aag aat tgg ttt gtt ggc ctc aag | 432 |
| Tyr Ile Ser Lys Lys His Ala Glu Lys Asn Trp Phe Val Gly Leu Lys |     |
| 130 135 140   |     |

aag aat ggg agc tgc aaa cgc ggt cct cgg act cac tat ggc cag aaa 480  
 Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg Thr His Tyr Gly Gln Lys  
 145 150 155 160

gca atc ttg ttt ctc ccc ctg cca gtc tct tct gat 516  
 Ala Ile Leu Phe Leu Pro Leu Pro Val Ser Ser Asp  
 165 170

<210> 27

<211> 210

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
 sequence for a part of mouse fibroblast growth factor 6 and  
 a part of human fibroblast growth 1

<400> 27

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val  
 1 5 10 15

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala  
 20 25 30

Arg Ala Asn ~~Gly~~ Thr Leu Leu Asp Ser Arg Gly Trp Gly Thr Leu Leu  
 35 40 45

Ser Arg Ser Arg Ala Gly Leu Ala Gly Glu Ile Ser Gly Val Asn Trp

50                      55                      60  
 Glu Ser Gly Tyr Leu Val Gly Ile Lys Arg Gln Ala Asn Tyr Lys Lys  
 65                      70                      75                      80  
 Pro Lys Leu Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu  
 85                      90                      95  
 Pro Asp Gly Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile  
 100                      105                      110  
 Gln Leu Gln Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser  
 115                      120                      125  
 Thr Glu Thr Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr  
 130                      135                      140  
 Gly Ser Gln Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu  
 145                      150                      155                      160  
 Glu Asn His Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn  
 165                      170                      175  
 Trp Phe Val Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg  
 180                      185                      190  
 Thr His Tyr Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser  
 195                      200                      205  
 Ser Asp  
 210

human FGf-1

<210> 28

<211> 630

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of  
sequence for a part of mouse fibroblast growth factor 6 and  
a part of human fibroblast growth 1

<220>

<221> CDS

<222> (1)..(630)

<400> 28

atg tcc cgg gga gca gga cgt gtt cag ggc acg ctg cag gct ctc gtc 48

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1 5 10 15

ttc tta ggc gtc cta gtg ggc atg gtg gtg ccc tca cct gcc ggc gcc 96

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20 25 30

cgc gcc aac ggc acg cta ctg gac tcc aga ggc tgg ggc acc ctc ttg 144

Arg Ala Asn Gly Thr Leu Leu Asp Ser Arg Gly Trp Gly Thr Leu Leu

35 40 45

tcc agg tct cga gct ggg cta gct gga gag att tcg ggt gtg aat tgg 192



|   |     |
|---|-----|
| Ser Arg Ser Arg Ala Gly Leu Ala Gly Glu Ile Ser Gly Val Asn Trp |     |
| 50 55 60  |     |
| gaa agc ggc tat ttg gtg ggc att aag cga cag gct aat tac aag aag | 240 |
| Glu Ser Gly Tyr Leu Val Gly Ile Lys Arg Gln Ala Asn Tyr Lys Lys |     |
| 65 70 75 80   |     |
| ccc aaa ctc ctc tac tgt agc aac ggg ggc cac ttc ctg agg atc ctt | 288 |
| Pro Lys Leu Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu |     |
| 85 90 95  |     |
| ccg gat ggc aca gtg gat ggg aca agg gac agg agc gac cag cac att | 336 |
| Pro Asp Gly Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile |     |
| 100 105 110   |     |
| cag ctg cag ctc agt gcg gaa agc gtg ggg gag gtg tat ata aag agt | 384 |
| Gln Leu Gln Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser |     |
| 115 120 125   |     |
| acc gag act ggc cag tac ttg gcc atg gac acc gac ggg ctt tta tac | 432 |
| Thr Glu Thr Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr |     |
| 130 135 140   |     |
| ggc tca cag aca cca aat gag gaa tgt ttg ttc ctg gaa agg ctg gag | 480 |
| Gly Ser Gln Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu |     |
| 145 150 155 160   |     |
| gag aac cat tac aac acc tat ata tcc aag aag cat gca gag aag aat | 528 |
| Glu Asn His Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu Lys Asn |     |
| 165 170 175   |     |

tgg ttt gtt ggc ctc aag aag aat ggg agc tgc aaa cgc ggt cct cgg 576  
 Trp Phe Val Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly Pro Arg

180 185 190

act cac tat ggc cag aaa gca atc ttg ttt ctc ccc ctg cca gtc tct 624  
 Thr His Tyr Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro Val Ser

195 200 205

tct gat 630

Ser Asp

210

<210> 29

<211> 180

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of mouse fibroblast growth factor 6,

a part of human fibroblast growth factor 1 and an artificial

sequence

<400> 29

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1 5 10 15

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20 25 30

human FGF-1

Arg Ala Asn Gly Thr Leu Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu

35

40

45

Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly

50

55

60

Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln

65

70

75

80

Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr

85

90

95

Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln

100

105

110

Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn Ala

115

120

125

Thr Pro Ala Pro His Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu

130

135

140

Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly

145

150

155

160

Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro

165

170

175

Val Ser Ser Asp

180

<210> 30

<211> 540

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: fusion of

sequence for a part of mouse fibroblast growth factor 6,  
a part of human fibroblast growth factor 1 and an artificial  
sequence

<220>

<221> CDS

<222> (1)..(540)

<400> 30

atg tcc cgg gga gca gga cgt gtt cag ggc acg ctg cag gct ctc gtc 48

Met Ser Arg Gly Ala Gly Arg Val Gln Gly Thr Leu Gln Ala Leu Val

1 5 10 15

ttc tta ggc gtc cta gtg ggc atg gtg gtg ccc tca cct gcc ggc gcc 96

Phe Leu Gly Val Leu Val Gly Met Val Val Pro Ser Pro Ala Gly Ala

20 25 30

cgc gcc aac ggc acg cta ctg gac gct aat tac aag aag ccc aaa ctc 144

Arg Ala Asn Gly Thr Leu Leu Asp Ala Asn Tyr Lys Lys Pro Lys Leu

35 40 45

ctc tac tgt agc aac ggg ggc cac ttc ctg agg atc ctt ccg gat ggc 192

|   |     |
|---|-----|
| Leu Tyr Cys Ser Asn Gly Gly His Phe Leu Arg Ile Leu Pro Asp Gly |     |
| 50 55 60  |     |
| aca gtg gat ggg aca agg gac agg agc gac cag cac att cag ctg cag | 240 |
| Thr Val Asp Gly Thr Arg Asp Arg Ser Asp Gln His Ile Gln Leu Gln |     |
| 65 70 75 80   |     |
| ctc agt gcg gaa agc gtg ggg gag gtg tat ata aag agt acc gag act | 288 |
| Leu Ser Ala Glu Ser Val Gly Glu Val Tyr Ile Lys Ser Thr Glu Thr |     |
| 85 90 95  |     |
| ggc cag tac ttg gcc atg gac acc gac ggg ctt tta tac ggc tca cag | 336 |
| Gly Gln Tyr Leu Ala Met Asp Thr Asp Gly Leu Leu Tyr Gly Ser Gln |     |
| 100 105 110   |     |
| aca cca aat gag gaa tgt ttg ttc ctg gaa agg ctg gag gag aac gct | 384 |
| Thr Pro Asn Glu Glu Cys Leu Phe Leu Glu Arg Leu Glu Glu Asn Ala |     |
| 115 120 125   |     |
| act cca gct cca cat tac aac acc tat ata tcc aag aag cat gca gag | 432 |
| Thr Pro Ala Pro His Tyr Asn Thr Tyr Ile Ser Lys Lys His Ala Glu |     |
| 130 135 140   |     |
| aag aat tgg ttt gtt ggc ctc aag aag aat ggg agc tgc aaa cgc ggt | 480 |
| Lys Asn Trp Phe Val Gly Leu Lys Lys Asn Gly Ser Cys Lys Arg Gly |     |
| 145 150 155 160   |     |
| cct cgg act cac tat ggc cag aaa gca atc ttg ttt ctc ccc ctg cca | 528 |
| Pro Arg Thr His Tyr Gly Gln Lys Ala Ile Leu Phe Leu Pro Leu Pro |     |
| 165 170 175   |     |

gtc tct tct gat

540

Val Ser Ser Asp

180

<210> 31

<211> 20

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:primer for PCR

<400> 31

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20

*β' included*